

Emergence of Rules in Cell Society: Differentiation, Hierarchy, and Stability

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Abstract

A dynamic model for cell differentiation is studied, where cells with internal chemical reaction dynamics interact with each other and replicate. It leads to spontaneous differentiation of cells and determination, as is discussed in the isologous diversification. Following features of the differentiation are obtained: (1) Hierarchical differentiation from a "stem" cell to other cell types, with the emergence of the interaction-dependent rules for differentiation; (2) Global stability of an ensemble of cells consisting of several cell types, that is sustained by the emergent, autonomous control on the rate of differentiation; (3) Existence of several cell colonies with different cell-type distributions. The results provide a novel viewpoint on the origin of complex cell society, while relevance to some biological problems, especially to the hemopoietic system, is also discussed.

1 Introduction

A multicellular organism is an ordered clone of a fertilized egg. All the cells contain the same genome set but are specialized in different ways. The emergence of different cell types is determined rather precisely, while the developmental process of the cells, viewed as a cell society, has robustness against perturbations.

In molecular biology, the differentiation processes are often regarded as on-off switching processes. Switch depends on inputs by signal molecules, which leads to a variety of cell types as outputs. A large number of reactions between inputs and outputs are represented as a "cascade", where the reactions are assumed to be approximately independent of the other reaction processes in the cell. The switching behavior (given by the sigmoidal function) is assumed to be generated from a chain of these reactions. With this viewpoint, one can decompose the differentiation by successive local elementary processes. It enables us to elucidate the differentiation processes by experimental methods, where several signal molecules and essential genes for differentiations are identified.

Of course, the development progresses through cooperation of several processes. Successive differentiation processes are often expressed as "canalization", where differentiations are captured as a result of dynamics of complex

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